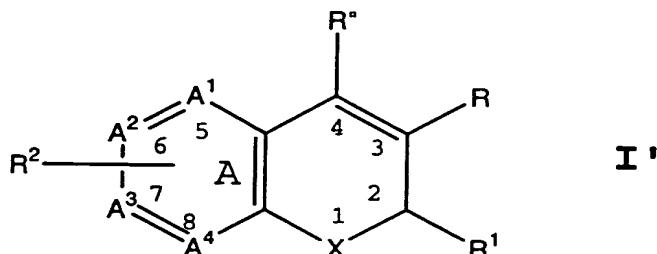


What is claimed is:

1. A compound of Formula I'

5



- wherein X is selected from O, S, CR^cR^b and NR^a;
- wherein R^a is selected from hydrido, C₁-C₃-alkyl,
10 (optionally substituted phenyl)-C₁-C₃-alkyl, acyl
and carboxy-C₁-C₆-alkyl;
- wherein each of R^b and R^c is independently selected
from hydrido, C₁-C₃-alkyl, phenyl-C₁-C₃-alkyl, C₁-
C₆-perfluoroalkyl, chloro, C₁-C₆-alkylthio, C₁-C₆-
alkoxy, nitro, cyano and cyano-C₁-C₃-alkyl;
- 15 wherein R is selected from carboxyl, aminocarbonyl,
C₁-C₆-alkylsulfonylaminocarbonyl and C₁-C₆-
alkoxycarbonyl;
- wherein R" is selected from hydrido, phenyl, thiienyl
and C₂-C₆-alkenyl;
- 20 wherein R¹ is selected from C₁-C₃-perfluoroalkyl,
chloro, C₁-C₆-alkylthio, C₁-C₆-alkoxy, nitro,
cyano and cyano-C₁-C₃-alkyl;
- wherein R² is one or more radicals independently
selected from hydrido, halo, C₁-C₆-alkyl, C₂-C₆-
25 alkenyl, C₂-C₆-alkynyl, halo-C₂-C₆-alkynyl, aryl-
C₁-C₆-alkyl, aryl-C₂-C₆-alkynyl, aryl-C₂-C₆-
alkenyl, C₁-C₆-alkoxy, methylenedioxy, C₁-C₆-
alkylthio, C₁-C₆-alkylsulfinyl, aryloxy,
arylothio, arylsulfinyl, heteroaryloxy, C₁-C₆-
30 alkoxy-C₁-C₆-alkyl, aryl-C₁-C₆-alkyloxy,
heteroaryl-C₁-C₆-alkyloxy, aryl-C₁-C₆-alkoxy-C₁-C₆-
alkyl, C₁-C₆-haloalkyl, C₁-C₆-haloalkoxy, C₁-C₆-
haloalkylthio, C₁-C₆-haloalkylsulfinyl, C₁-C₆-

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haloalkylsulfonyl, $C_1\text{-}C_6\text{-}($ haloalkyl- $C_1\text{-}C_6\text{-}$
 hydroxyalkyl, $C_1\text{-}C_6\text{-hydroxyalkyl}$, hydroxyimino- $C_1\text{-}$
 $C_6\text{-alkyl}$, $C_1\text{-}C_6\text{-alkylamino}$, arylamino, aryl- $C_1\text{-}C_6\text{-}$
 alkylamino, heteroaryl amino, heteroaryl- $C_1\text{-}C_6\text{-}$
 5 alkylamino, nitro, cyano, amino, aminosulfonyl,
 $C_1\text{-}C_6\text{-alkylaminosulfonyl}$, arylaminosulfonyl,
 heteroarylaminosulfonyl, aryl- $C_1\text{-}C_6\text{-}$
 alkylaminosulfonyl, heteroaryl- $C_1\text{-}C_6\text{-}$
 alkylaminosulfonyl, heterocyclsulfonyl, $C_1\text{-}C_6\text{-}$
 10 alkylsulfonyl, aryl- $C_1\text{-}C_6\text{-alkylsulfonyl}$,
 optionally substituted aryl, optionally
 substituted heteroaryl, aryl- $C_1\text{-}C_6\text{-alkylcarbonyl}$,
 heteroaryl- $C_1\text{-}C_6\text{-alkylcarbonyl}$,
 heteroarylcarbonyl, arylcarbonyl, aminocarbonyl,
 15 $C_1\text{-}C_6\text{-alkoxycarbonyl}$, formyl, $C_1\text{-}C_6\text{-}$
 haloalkylcarbonyl and $C_1\text{-}C_6\text{-alkylcarbonyl}$; and
 wherein the A ring atoms A^1 , A^2 , A^3 and A^4 are
 independently selected from carbon and nitrogen
 with the proviso that at least two of A^1 , A^2 , A^3
 20 and A^4 are carbon;
 or wherein R^2 together with ring A forms a radical
 selected from naphthyl, quinolyl, isoquinolyl,
 quinolizinyl, quinoxaliny and dibenzofuryl;
 or an isomer or pharmaceutically acceptable salt
 25 thereof.

2. A compound of Claim 1 wherein X is selected
 from O, S, CR^aR^b and NR^a ; wherein R^a is selected from
 hydrido, $C_1\text{-}C_6\text{-alkyl}$, (optionally substituted
 30 phenyl)- $C_1\text{-}C_6\text{-alkyl}$, acyl and carboxy- $C_1\text{-}C_6\text{-alkyl}$;
 wherein each of R^a and R^b is independently selected
 from hydrido, $C_1\text{-}C_6\text{-alkyl}$, phenyl- $C_1\text{-}C_6\text{-alkyl}$, $C_1\text{-}C_6\text{-}$
 perfluoroalkyl, chloro, $C_1\text{-}C_6\text{-alkylthio}$, $C_1\text{-}C_6\text{-alkoxy}$,
 nitro, cyano and cyano- $C_1\text{-}C_6\text{-alkyl}$; wherein R is
 35 selected from carboxyl, aminocarbonyl, $C_1\text{-}C_6\text{-}$
 alkylsulfonylaminocarbonyl and $C_1\text{-}C_6\text{-alkoxycarbonyl}$;
 wherein R^a is selected from hydrido, phenyl, thiényl
 and $C_2\text{-}C_6\text{-alkenyl}$; wherein R^b is selected from $C_1\text{-}C_6\text{-}$

perfluoroalkyl, chloro, C₁-C₆-alkylthio, C₁-C₆-alkoxy, nitro, cyano and cyano-C₁-C₆-alkyl; wherein R² is one or more radicals independently selected from hydrido, halo, C₁-C₆-alkyl, C₂-C₆-alkenyl, C₂-C₆-alkynyl, halo-C₂-C₆-alkynyl, aryl-C₁-C₆-alkyl, aryl-C₂-C₆-alkynyl, aryl-C₂-C₆-alkenyl, C₁-C₆-alkoxy, methylenedioxy, C₁-C₆-alkylthio, C₁-C₆-alkylsulfinyl, aryloxy, arylthio, arylsulfinyl, heteroaryloxy, C₁-C₆-alkoxy-C₁-C₆-alkyl, aryl-C₁-C₆-alkyloxy, heteroaryl-C₁-C₆-alkyloxy, aryl-C₁-C₆-alkoxy-C₁-C₆-alkyl, C₁-C₆-haloalkyl, C₁-C₆-haloalkoxy, C₁-C₆-haloalkylthio, C₁-C₆-haloalkylsulfinyl, C₁-C₆-haloalkylsulfonyl, C₁-C₆-(haloalkyl-C₁-C₆-hydroxyalkyl, C₁-C₆-hydroxyalkyl, hydroxyimino-C₁-C₆-alkyl, C₁-C₆-alkylamino, arylamino, aryl-C₁-C₆-alkylamino, heteroaryl-amino, heteroaryl-C₁-C₆-alkylamino, nitro, cyano, amino, aminosulfonyl, C₁-C₆-alkylaminosulfonyl, arylaminosulfonyl, heteroarylaminosulfonyl, aryl-C₁-C₆-alkylaminosulfonyl, heteroaryl-C₁-C₆-alkylaminosulfonyl, heteroaryl-C₁-C₆-alkylcarbonyl, heterocyclsulfonyl, C₁-C₆-alkylsulfonyl, aryl-C₁-C₆-alkylsulfonyl, optionally substituted aryl, optionally substituted heteroaryl, aryl-C₁-C₆-alkylcarbonyl, heteroaryl-C₁-C₆-alkylcarbonyl, heteroarylcarbonyl, arylcarbonyl, aminocarbonyl, C₁-C₆-alkoxycarbonyl, formyl, C₁-C₆-haloalkylcarbonyl and C₁-C₆-alkylcarbonyl; and wherein the A ring atoms A¹, A², A³ and A⁴ are independently selected from carbon and nitrogen with the proviso that at least three of A¹, A², A³ and A⁴ are carbon; or wherein R² together with ring A forms a naphthyl or quinolyl radical; or an isomer or pharmaceutically acceptable salt thereof.

3. A compound of Claim 2 wherein X is selected from O, S and NR⁴; wherein R⁴ is selected from hydrido, C₁-C₆-alkyl and (optionally substituted phenyl)methyl; wherein R is carboxyl; wherein R⁵ is selected from hydrido and C₂-C₆-alkenyl; wherein R¹ is

selected from C₁-C₆-perfluoroalkyl; wherein R² is one or more radicals independently selected from hydrido, halo, C₁-C₆-alkyl, C₂-C₆-alkenyl, C₂-C₆-alkynyl, halo-C₁-C₆-alkynyl, phenyl-C₁-C₆-alkyl,
5 phenyl-C₂-C₆-alkynyl, phenyl-C₂-C₆-alkenyl, C₁-C₆-alkoxy, methylenedioxy, C₁-C₃-alkoxy-C₁-C₃-alkyl, C₁-C₃-alkylthio, C₁-C₃-alkylsulfinyl, phenyloxy,
phenylthio, phenylsulfinyl, C₁-C₃-haloalkyl-C₁-C₃-hydroxyalkyl, phenyl-C₁-C₃-alkyloxy-C₁-C₃-alkyl, C₁-C₆-
10 haloalkyl, C₁-C₆-haloalkoxy, C₁-C₆-haloalkylthio, C₁-C₆-hydroxyalkyl, C₁-C₆-alkoxy-C₁-C₃-alkyl,
hydroxyimino-C₁-C₆-alkyl, C₁-C₆-alkylamino, nitro,
cyano, amino, aminosulfonyl, N-alkylaminosulfonyl,
N-arylaminosulfonyl, N-heteroarylaminosulfonyl, N-
15 (phenyl-C₁-C₆-alkyl)aminosulfonyl, N-(heteroaryl-C₁-C₆-alkyl)aminosulfonyl, phenyl-C₁-C₆-alkylsulfonyl, 5-
to 8-membered heterocyclsulfonyl, C₁-C₆-alkylsulfonyl, optionally substituted phenyl,
optionally substituted 5- to 9-membered heteroaryl,
20 phenyl-C₁-C₆-alkylcarbonyl, phenylcarbonyl, 4-chlorophenylcarbonyl, 4-hydroxyphenylcarbonyl, 4-trifluoromethylphenylcarbonyl, 4-methoxyphenylcarbonyl, aminocarbonyl, formyl, and C₁-C₆-alkylcarbonyl; wherein the A ring atoms A¹, A², A³
25 and A⁴ are independently selected from carbon and nitrogen with the proviso that at least three of A¹, A², A³ and A⁴ are carbon; or wherein R² together with ring A forms a naphthyl, benzofurylphenyl, or quinolyl radical; or an isomer or pharmaceutically
30 acceptable salt thereof.

4. A compound of Claim 3 wherein X is selected from O, S and NR^a; wherein R^a is selected from hydrido, methyl, ethyl, (4-trifluoromethyl)benzyl,
35 (4-chloromethyl)benzyl, (4-methoxy)benzyl, and (4-cyano)benzyl, (4-nitro)benzyl; wherein R is carboxyl; wherein R" is selected from hydrido and ethenyl; wherein R¹ is selected from trifluoromethyl

- and pentafluoroethyl; wherein R² is one or more radicals independently selected from hydrido, chloro, bromo, fluoro, iodo, methyl, tert-butyl, ethenyl, ethynyl, 5-chloro-1-pentynyl, 1-pentynyl,
- 5 3,3-dimethyl-1-butynyl, benzyl, phenylethyl, phenyl-ethynyl, 4-chlorophenyl-ethynyl, 4-methoxyphenyl-ethynyl, phenylethenyl, methoxy, methylthio, methylsulfinyl, phenoxy, phenylthio, phenylsulfinyl, methylenedioxy, benzyloxymethyl,
- 10 trifluoromethyl, difluoromethyl, pentafluoroethyl, trifluoromethoxy, trifluoromethylthio, hydroxymethyl, hydroxy-trifluoroethyl, methoxymethyl, hydroxyiminomethyl, N-methylamino, nitro, cyano, amino, aminosulfonyl, N-
- 15 methylaminosulfonyl, N-phenylaminosulfonyl, N-furylaminosulfonyl, N-(benzyl)aminosulfonyl, N-(furylmethyl)aminosulfonyl, benzylsulfonyl, phenylethylaminosulfonyl, furylsulfonyl, methylsulfonyl, phenyl, phenyl substituted with one
- 20 or more radicals selected from chloro, fluoro, bromo, methoxy, methylthio and methylsulfonyl, benzimidazolyl, thienyl, thienyl substituted with chloro, furyl, furyl substituted with chloro, benzylcarbonyl, optionally substituted
- 25 phenylcarbonyl, aminocarbonyl, formyl and methylcarbonyl; wherein the A ring atoms A¹, A², A³ and A⁴ are independently selected from carbon and nitrogen with the proviso that at least three of A¹, A², A³ and A⁴ are carbon; or wherein R² together with
- 30 ring A forms a naphthyl, or quinolyl radical; or an isomer or pharmaceutically acceptable salt thereof.

5. A compound of Claim 4 selected from compounds, and their isomers and pharmaceutically-
35 acceptable salts, of the group consisting of 6-chloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;

- 7-ethyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 7-methyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 5 2,7-bis(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 7-bromo-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-chloro-7-methyl-2-trifluoromethyl-2H-1-benzopyran-
- 10 3-carboxylic acid;
- 8-(1-methylethyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-chloro-7-(1,1-dimethylethyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 15 6-chloro-8-(1-methylethyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 8-ethoxy-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 20 7-(1,1-dimethylethyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-bromo-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 8-chloro-2-trifluoromethyl-2H-1-benzopyran-3-
- 25 carboxylic acid;
- 8-bromo-6-chloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-trifluoromethoxy-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 30 8-fluoro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 5,7-dichloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 7,8-dichloro-2-trifluoromethyl-2H-1-benzopyran-3-
- 35 carboxylic acid;
- 7-isopropoxy-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;

- 8-phenyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 7,8-dimethyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 5 6,8-bis(1,1-dimethylethyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 7-chloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 7-(1-methylethyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 10 7-phenyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 7-phenyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-chloro-7-ethyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 15 8-ethyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-chloro-8-ethyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-chloro-7-phenyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 20 6,7-dichloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6,8-dichloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 25 6,8-dibromo-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6,8-dimethoxy-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-nitro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 30 6-amino-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- ethyl 6-amino-2-trifluoromethyl-2H-1-benzopyran-3-carboxylate;
- 35 6-chloro-8-methyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 8-chloro-6-methyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;

- 8-chloro-6-methoxy-2-trifluoromethyl-2H-1-
benzopyran-3-carboxylic acid;
- 6,8-difluoro-2-trifluoromethyl-2H-1-benzopyran-3-
carboxylic acid;
- 5 6-bromo-8-chloro-2-trifluoromethyl-2H-1-benzopyran-
3-carboxylic acid;
- 8-bromo-6-fluoro-2-trifluoromethyl-2H-1-benzopyran-
3-carboxylic acid;
- 8-bromo-6-methyl-2-trifluoromethyl-2H-1-benzopyran-
10 3-carboxylic acid;
- 8-bromo-5-fluoro-2-trifluoromethyl-2H-1-benzopyran-
3-carboxylic acid;
- 6-chloro-8-fluoro-2-trifluoromethyl-2H-1-benzopyran-
3-carboxylic acid;
- 15 6-bromo-8-methoxy-2-trifluoromethyl-2H-1-benzopyran-
3-carboxylic acid;
- 7-(N,N-diethylamino)-2-trifluoromethyl-2H-1-
benzopyran-3-carboxylic acid;
- 6-[(phenylmethyl)amino]sulfonyl]-2-trifluoromethyl-
20 2H-1-benzopyran-3-carboxylic acid;
- 6-[(dimethylamino)sulfonyl]-2-trifluoromethyl-2H-1-
benzopyran-3-carboxylic acid;
- 6-aminosulfonyl-2-trifluoromethyl-2H-1-benzopyran-3-
carboxylic acid;
- 25 6-(methylamino)sulfonyl-2-trifluoromethyl-2H-1-
benzopyran-3-carboxylic acid;
- 6-[(4-morpholino)sulfonyl]-2-trifluoromethyl-2H-1-
benzopyran-3-carboxylic acid;
- 6-[(1,1-dimethylethyl)aminosulfonyl]-2-
30 trifluoromethyl-2H-1-benzopyran-3-carboxylic
acid;
- 6-[(2-methylpropyl)aminosulfonyl]-2-trifluoromethyl-
2H-1-benzopyran-3-carboxylic acid;
- 6-methylsulfonyl-2-trifluoromethyl-2H-1-benzopyran-
35 3-carboxylic acid;
- 8-chloro-6-[(phenylmethyl)amino]sulfonyl]-2-
trifluoromethyl-2H-1-benzopyran-3-carboxylic
acid;

- 6-N,N-diethylaminosulfonyl-2-trifluoromethyl-2H-1-
benzopyran-3-carboxylic acid;
- 6-phenylacetyl-2-trifluoromethyl-2H-1-benzopyran-3-
carboxylic acid;
- 5 6-(2,2-dimethylpropylcarbonyl)-2-trifluoromethyl-2H-
1-benzopyran-3-carboxylic acid;
- 6,8-dichloro-7-methoxy-2-trifluoromethyl-2H-1-
benzopyran-3-carboxylic acid;
- 6-chloro-2-trifluoromethyl-2H-1-benzothiopyran-3-
10 carboxylic acid;
- 6-[(2-furanylmethyl)amino]sulfonyl]-2-
(trifluoromethyl)-2H-1-benzopyran-3-carboxylic
acid;
- 6-[(phenylmethyl)sulfonyl]-2-(trifluoromethyl)-2H-1-
15 benzopyran-3-carboxylic acid;
- 6-[(phenylethyl)amino]sulfonyl]-2-
(trifluoromethyl)-2H-1-benzopyran-3-carboxylic
acid;
- 6-iodo-2-trifluoromethyl-2H-1-benzopyran-3-
20 carboxylic acid;
- 6-chloro-8-ido-2-(trifluoromethyl)-2H-1-benzopyran-
3-carboxylic acid;
- 8-bromo-6-chloro-2-trifluoromethyl-2H-1-benzopyran-
3-carboxylic acid;
- 25 6-formyl-2-(trifluoromethyl)-2H-1-benzopyran-3-
carboxylic acid;
- 6-chloro-8-formyl-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;
- 6-bromo-7-(1,1-dimethylethyl)-2-(trifluoromethyl)-
30 2H-1-benzopyran-3-carboxylic acid;
- 5,6-dichloro-2-(trifluoromethyl)-2H-1-benzopyran-3-
carboxylic acid;
- 6-cyano-2-(trifluoromethyl)-2H-1-benzopyran-3-
carboxylic acid ;
- 35 6-hydroxymethyl-2-(trifluoromethyl)-2H-1-benzopyran-
3-carboxylic acid;
- 6-(difluoromethyl)-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;

- 2,6-bis(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 5,6,7-trichloro-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 5 6,7,8-trichloro-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-(methylthio)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-(methylsulfinyl)-2-(trifluoromethyl)-2H-1-
- 10 benzopyran-3-carboxylic acid;
- 5,8-dichloro-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-(pentafluoroethyl)-2-(trifluoromethyl)-2H-1-
- benzopyran-3-carboxylic acid;
- 15 6-(1,1-dimethylethyl)-2-(trifluoromethyl)-2H-1-
- benzopyran-3-carboxylic acid;
- 2-(trifluoromethyl)-6-[(trifluoromethyl)thio]-2H-1-
- benzopyran-3-carboxylic acid;
- 6,8-dichloro-7-methyl-2-(trifluoromethyl)-2H-1-
- 20 benzopyran-3-carboxylic acid;
- 6-chloro-2,7-bis(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 5-methoxy-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 25 6-benzoyl-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-(4-chlorobenzoyl)-2-(trifluoromethyl)-2H-1-
- benzopyran-3-carboxylic acid;
- 6-(4-hydroxybenzoyl)-2-(trifluoromethyl)-2H-1-
- 30 benzopyran-3-carboxylic acid;
- 6-phenoxy-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 8-chloro-6-(4-chlorophenoxy)-2-(trifluoromethyl)-2H-
- 1-benzopyran-3-carboxylic acid;
- 35 2-(trifluoromethyl)-6-[4-(trifluoromethyl)phenoxy]-2H-1-benzopyran-3-carboxylic acid;
- 6-(4-methoxyphenoxy)-2-(trifluoromethyl)-2H-1-
- benzopyran-3-carboxylic acid;

- 6-(3-chloro-4-methoxyphenoxy)-2-(trifluoromethyl)-
2H-1-benzopyran-3-carboxylic acid;
- 6-(4-chlorophenoxy)-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;
- 5 8-chloro-2-(trifluoromethyl)-6-[4-
(trifluoromethyl)phenoxy]-2H-1-benzopyran-3-
carboxylic acid;
- 6-chloro-8-cyano-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;
- 10 6-chloro-8-[(hydroxyimino)methyl]-2-
(trifluoromethyl)-2H-1-benzopyran-3-carboxylic
acid;
- 6-chloro-8-(hydroxymethyl)-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;
- 15 8-(1H-benzimidazol-2-yl)-6-chloro-2-
(trifluoromethyl)-2H-1-benzopyran-3-carboxylic
acid;
- 7-(1,1-dimethylethyl)-2-(pentafluoroethyl)-2H-1-
benzopyran-3-carboxylic acid;
- 20 6-chloro-8-(methoxymethyl)-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;
- 6-chloro-8-(benzyloxymethyl)-2-(trifluoromethyl)-2H-
1-benzopyran-3-carboxylic acid;
- 6-chloro-8-ethenyl-2-(trifluoromethyl)-2H-1-
25 benzopyran-3-carboxylic acid;
- 6-chloro-8-ethynyl-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;
- 6-chloro-8-(2-thienyl)-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;
- 30 6-chloro-8-(2-furanyl)-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;
- 6-chloro-8-(5-chloro-1-pentynyl)-2-
(trifluoromethyl)-2H-1-benzopyran-3-carboxylic
acid;
- 35 6-chloro-8-(1-pentynyl)-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;
- 6-chloro-8-(phenylethylyn)-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;

6-chloro-8-(3,3-dimethyl-1-butynyl)-2-
(trifluoromethyl)-2H-1-benzopyran-3-carboxylic
acid;

6-chloro-8-[(4-chlorophenyl)ethynyl]-2-
5 (trifluoromethyl)-2H-1-benzopyran-3-carboxylic
acid;

6-chloro-8-[(4-methoxyphenyl)ethynyl]-2-
(trifluoromethyl)-2H-1-benzopyran-3-carboxylic
acid;

10 6-(phenylethynyl)-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;

6-chloro-8-(4-chlorophenyl)-2-(trifluoromethyl)-2H-
1-benzopyran-3-carboxylic acid;

6-chloro-8-(3-methoxyphenyl)-2-(trifluoromethyl)-2H-
15 1-benzopyran-3-carboxylic acid;

6-chloro-8-[(4-methylthio)phenyl]-2-
(trifluoromethyl)-2H-1-benzopyran-3-carboxylic
acid;

6-chloro-8-[(4-methylsulfonyl)phenyl]-2-

20 (trifluoromethyl)-2H-1-benzopyran-3-carboxylic
acid;

6-chloro-8-phenyl-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;

6-bromo-8-fluoro-2-(trifluoromethyl)-2H-1-
25 benzopyran-3-carboxylic acid;

6-(4-fluorophenyl)-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;

6-phenyl-2-(trifluoromethyl)-2H-1-benzopyran-3-
carboxylic acid;

30 8-chloro-6-fluoro-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;

6,8-diido-2-(trifluoromethyl)-2H-1-benzopyran-3-
carboxylic acid;

6-(5-chloro-2-thienyl)-2-(trifluoromethyl)-2H-1-
35 benzopyran-3-carboxylic acid;

6-(2-thienyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-
carboxylic acid;

- 6-(4-chlorophenyl)-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;
- 6-(4-bromophenyl)-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;
- 5 6-(ethynyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-
carboxylic acid;
- 6-methyl-2-(trifluoromethyl)-2H-1-benzopyran-3-
carboxylic acid;
- 6-chloro-8-(4-methoxyphenyl)-2-trifluoromethyl-2H-1-
10 benzopyran-3-carboxylic acid;
- 6-chloro-2-(trifluoromethyl)-4-ethenyl-2H-1-
benzopyran-3-carboxylic acid;
- 6-chloro-2-(trifluoromethyl)-4-phenyl-2H-1-
benzopyran-3-carboxylic acid;
- 15 6-chloro-4-(2-thienyl)-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;
- 6-(2,2,2-trifluoro-1-hydroxyethyl)-2-
(trifluoromethyl)-2H-1-benzopyran-3-carboxylic
acid;
- 20 6-methyl-2-(trifluoromethyl)-2H-1-benzothiopyran-3-
carboxylic acid;
- 6,8-dimethyl-2-(trifluoromethyl)-2H-1-
benzothiopyran-3-carboxylic acid;
- 25 6-(1,1-dimethylethyl)-2-(trifluoromethyl)-2H-1-
benzothiopyran-3-carboxylic acid;
- 7-methyl-2-(trifluoromethyl)-2H-1-benzothiopyran-3-
carboxylic acid;
- 6,7-dimethyl-2-(trifluoromethyl)-2H-1-
30 benzothiopyran-3-carboxylic acid;
- 8-methyl-2-(trifluoromethyl)-2H-1-benzothiopyran-3-
carboxylic acid;
- 2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic
acid;
- 35 6-chloro-7-methyl-2-(trifluoromethyl)-2H-1-
benzothiopyran-3-carboxylic acid;
- 7-chloro-2-(trifluoromethyl)-2H-1-benzothiopyran-3-
carboxylic acid;

- 6,7-dichloro-2-(trifluoromethyl)-2H-1-
benzothiopyran-3-carboxylic acid;
2-(trifluoromethyl)-6-[(trifluoromethyl)thio]-2H-1-
benzopyran-3-carboxylic acid;
- 5 6,8-dichloro-2-trifluoromethyl-2H-1-benzothiopyran-
3-carboxylic acid;
6-chloro-1,2-dihydro-2-(trifluoromethyl)-3-
quinolinecarboxylic acid;
6,8-dichloro-1,2-dihydro-2-(trifluoromethyl)-3-
10 quinolinecarboxylic acid;
6,7-difluoro-1,2-dihydro-2-(trifluoromethyl)-3-
quinolinecarboxylic acid;
6-iodo-1,2-dihydro-2-(trifluoromethyl)-3-
quinolinecarboxylic acid;
- 15 6-bromo-1,2-dihydro-2-(trifluoromethyl)-3-
quinolinecarboxylic acid;
1,2-dihydro-6-(trifluoromethoxy)-2-
(trifluoromethyl)-3-quinolinecarboxylic acid;
- 20 6-(trifluoromethyl)-1,2-dihydro-2-(trifluoromethyl)-
3-quinolinecarboxylic acid;
6-cyano-1,2-dihydro-2-(trifluoromethyl)-3-
quinolinecarboxylic acid;
- 25 6-chloro-1,2-dihydro-2-(trifluoromethyl)-1-[(4-
(trifluoromethyl)phenyl)methyl]-3-
quinolinecarboxylic acid;
- 6-chloro-1-[(4-chlorophenyl)methyl]-1,2-dihydro-2-
(trifluoromethyl)-3-quinolinecarboxylic acid;
- 30 35 6-chloro-1,2-dihydro-2-(trifluoromethyl)-1-[(4-
(methoxy)phenyl)methyl]-3-quinolinecarboxylic
acid;
6-chloro-1-[(4-cyanophenyl)methyl]-1,2-dihydro-2-
(trifluoromethyl)-3-quinolinecarboxylic acid;
- 6-chloro-1-[(4-nitrophenyl)methyl]-2-
(trifluoromethyl)-3-quinolinecarboxylic acid;
6-chloro-1,2-dihydro-1-ethyl-2-(trifluoromethyl)-3-
quinolinecarboxylic acid;

6-chloro-2-(trifluoromethyl)-1,2-dihydro[1,8]naphthyridine-3-carboxylic acid; 2-trifluoromethyl-2H-naphtho[1,2-b]pyran-3-carboxylic acid;

- 5 2-trifluoromethyl-3H-naphtho[2,1-b]pyran-3-carboxylic acid;
2-trifluoromethyl-2H-naphtho[2,3-b]pyran-3-carboxylic acid;
5-(hydroxymethyl)-8-methyl-2-(trifluoromethyl)-2H-pyrano[2,3-c]pyridine-3-carboxylic acid;
0 6-(trifluoromethyl)-6h-1,3-dioxolo[4,5-g][1]benzopyran-7-carboxylic acid; and
3-(trifluoromethyl)-3H-benzofuro[3,2-f][1]benzopyran-2-carboxylic acid.

15

6. A compound of Claim 2 wherein X is O; wherein R is carboxyl; wherein R" is selected from hydrido and C₂-C₆-alkenyl; wherein R¹ is selected from C₁-C₃-perfluoroalkyl; wherein R² is one or more radicals independently selected from hydrido, halo, C₁-C₆-alkyl, phenyl-C₁-C₆-alkyl, phenyl-C₂-C₆-alkynyl, phenyl-C₂-C₆-alkenyl, C₁-C₆-alkoxy, phenoxy, 5- or 6-membered heteroaryloxy, phenyl-C₁-C₆-alkyloxy, 5- or 6-membered heteroaryl-C₁-C₆-alkyloxy, C₁-C₆-haloalkyl, C₁-C₆-haloalkoxy, N-(C₁-C₆-alkyl)amino, N,N-di-(C₁-C₆-alkyl)amino, N-phenylamino, N-(phenyl-C₁-C₆-alkyl)amino, N-heteroarylamino, N-(heteroaryl-C₁-C₆-alkylamino, nitro, amino, aminosulfonyl, N-(C₁-C₆-alkyl)aminosulfonyl, N,N-di-(C₁-C₆-alkyl)aminosulfonyl, N-arylaminosulfonyl, N-heteroarylaminosulfonyl, N-(phenyl-C₁-C₆-alkyl)aminosulfonyl, N-(heteroaryl-C₁-C₆-alkyl)aminosulfonyl, 5- to 8-membered heterocyclsulfonyl, C₁-C₆-alkylsulfonyl, optionally substituted phenyl, optionally substituted 5- or 6-membered heteroaryl, phenyl-C₁-C₆-alkylcarbonyl, heteroarylcarbonyl, phenylcarbonyl, aminocarbonyl,

and C₁-C₆-alkylcarbonyl; wherein the A ring atoms A¹, A², A³ and A⁴ are independently selected from carbon and nitrogen with the proviso that at least three of A¹, A², A³ and A⁴ are carbon; or an isomer or
5 pharmaceutically acceptable salt thereof.

7. A compound of Claim 6 wherein X is O;
wherein R is carboxyl; wherein R" is selected from
hydrido and ethenyl; wherein R¹ is selected from
10 trifluoromethyl and pentafluoroethyl; wherein R² is
one or more radicals independently selected from
hydrido, chloro, bromo, fluoro, iodo, methyl, tert-
butyl, ethenyl, ethynyl, 5-chloro-1-pentynyl, 1-
pentynyl, 3,3-dimethyl-1-butynyl, benzyl,
15 phenylethyl, phenyl-ethynyl, 4-chlorophenyl-ethynyl,
4-methoxyphenyl-ethynyl, phenylethenyl, methoxy,
methylthio, methylsulfinyl, phenoxy, phenylthio,
phenylsulfinyl, pyridyloxy, thienyloxy, furyloxy,
phenylmethoxy, methylenedioxy, benzyloxymethyl,
20 trifluoromethyl, difluoromethyl, pentafluoroethyl,
trifluoromethoxy, trifluoromethylthio,
hydroxymethyl, hydroxy-trifluoroethyl,
methoxymethyl, hydroxyiminomethyl, N-methylamino, N-
phenylamino, N-(benzyl)amino, nitro, cyano, amino,
25 aminosulfonyl, N-methylaminosulfonyl, N-
phenylaminosulfonyl, N-furylaminosulfonyl, N-
(benzyl)aminosulfonyl, N-(furylmethyl)aminosulfonyl,
benzylsulfonyl, phenylethylaminosulfonyl,
furylsulfonyl, methylsulfonyl, phenyl, phenyl
30 substituted with one or more radicals selected from
chloro, fluoro, bromo, methoxy, methylthio and
methylsulfonyl, benzimidazolyl, thienyl, thietyl
substituted with chloro, furyl, furyl substituted
with chloro, benzylcarbonyl, furylcarbonyl,
35 phenylcarbonyl, aminocarbonyl, formyl, and
methylcarbonyl; and wherein one of the A ring atoms
A¹, A², A³ and A⁴ is nitrogen and the other three are

carbon; or an isomer or pharmaceutically acceptable salt thereof.

9. A compound of Claim 7 wherein X is O;
- 5 wherein R is carboxyl; wherein R" is selected from hydrido and ethenyl; wherein R¹ is selected from trifluoromethyl and pentafluoroethyl; wherein R² is one or more radicals independently selected from hydrido, chloro, bromo, fluoro, iodo, methyl, tert-10 butyl, ethenyl, ethynyl, 5-chloro-1-pentynyl, 1-pentynyl, 3,3-dimethyl-1-butynyl, benzyl, phenylethyl, phenyl-ethynyl, 4-chlorophenyl-ethynyl, 4-methoxyphenyl-ethynyl, phenylethenyl, methoxy, methylthio, methylsulfinyl, phenoxy, phenylthio, 15 phenylsulfinyl, pyridyloxy, thienyloxy, furyloxy, phenylmethoxy, methylenedioxy, benzyloxymethyl, trifluoromethyl, difluoromethyl, pentafluoroethyl, trifluoromethoxy, trifluoromethylthio, hydroxymethyl, hydroxy-trifluoroethyl, 20 methoxymethyl, hydroxyiminomethyl, N-methylamino, N-phenylamino, N-(benzyl)amino, nitro, cyano, amino, aminosulfonyl, N-methylaminosulfonyl, N-phenylaminosulfonyl, N-furylaminosulfonyl, N-(benzyl)aminosulfonyl, N-(furylmethyl)aminosulfonyl, 25 benzylsulfonyl, phenylethylaminosulfonyl, furylsulfonyl, methylsulfonyl, phenyl, phenyl substituted with one or more radicals selected from chloro, fluoro, bromo, methoxy, methylthio and methylsulfonyl, benzimidazolyl, thienyl, thienyl 30 substituted with chloro, furyl, furyl substituted with chloro, benzylcarbonyl, furylcarbonyl, phenylcarbonyl, aminocarbonyl, formyl, and methylcarbonyl; wherein the A ring atoms A¹, A², A³ and A⁴ are carbon; or an isomer or pharmaceutically 35 acceptable salt thereof.

10. A compound of Claim 9 selected from compounds, and their isomers and pharmaceutically-acceptable salts, of the group consisting of
6-chloro-2-trifluoromethyl-2H-1-benzopyran-3-
5 carboxylic acid;
(S)-6-chloro-2-trifluoromethyl-2H-1-benzopyran-3-
carboxylic acid;
6-chloro-7-methyl-2-trifluoromethyl-2H-1-benzopyran-
3-carboxylic acid;
10 6-chloro-7-(1,1-dimethylethyl)-2-trifluoromethyl-2H-
1-benzopyran-3-carboxylic acid;
(S)-6-chloro-7-(1,1-dimethylethyl)-2-
trifluoromethyl-2H-1-benzopyran-3-carboxylic
acid;
15 6-chloro-8-(1-methylethyl)-2-trifluoromethyl-2H-1-
benzopyran-3-carboxylic acid;
7-(1,1-dimethylethyl)-2-trifluoromethyl-2H-1-
benzopyran-3-carboxylic acid;
6-trifluoromethoxy-2-trifluoromethyl-2H-1-
20 benzopyran-3-carboxylic acid;
(S)-6-trifluoromethoxy-2-trifluoromethyl-2H-1-
benzopyran-3-carboxylic acid;
6,7-dichloro-2-trifluoromethyl-2H-1-benzopyran-3-
carboxylic acid;
25 6,8-dichloro-2-trifluoromethyl-2H-1-benzopyran-3-
carboxylic acid;
(S)-6,8-dichloro-2-trifluoromethyl-2H-1-benzopyran-
3-carboxylic acid;
6,8-dichloro-7-methoxy-2-trifluoromethyl-2H-1-
30 benzopyran-3-carboxylic acid;
6-chloro-2-trifluoromethyl-2H-1-benzothiopyran-3-
carboxylic acid;
(S)-6-chloro-2-trifluoromethyl-2H-1-benzothiopyran-
3-carboxylic acid;
35 6-cyano-2-(trifluoromethyl)-2H-1-benzopyran-3-
carboxylic acid;
(S)-6-cyano-2-(trifluoromethyl)-2H-1-benzopyran-3-
carboxylic acid;

- 6-hydroxymethyl-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-(difluoromethyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 5 2,6-bis(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 5,6,7-trichloro-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6,7,8-trichloro-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 10 6-(methylthio)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-(pentafluoroethyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 15 2-(trifluoromethyl)-6-[(trifluoromethyl)thio]-2H-1-benzopyran-3-carboxylic acid;
- 6,8-dichloro-7-methyl-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-benzoyl-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 20 6-(4-chlorobenzoyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-(4-hydroxybenzoyl)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 25 6-phenoxy-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 2-(trifluoromethyl)-6-[4-(trifluoromethyl)phenoxy]-2H-1-benzopyran-3-carboxylic acid;
- (S)-2-(trifluoromethyl)-6-[4-
- 30 (trifluoromethyl)phenoxy]-2H-1-benzopyran-3-carboxylic acid;
- 6-(4-methoxyphenoxy)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;
- 6-(3-chloro-4-methoxyphenoxy)-2-(trifluoromethyl)-
- 35 2H-1-benzopyran-3-carboxylic acid;
- 6-(4-chlorophenoxy)-2-(trifluoromethyl)-2H-1-benzopyran-3-carboxylic acid;

- 8-chloro-2-(trifluoromethyl)-6-[4-
(trifluoromethyl)phenoxy]-2H-1-benzopyran-3-
carboxylic acid;
- 6-chloro-8-cyano-2-(trifluoromethyl)-2H-1-
5 benzopyran-3-carboxylic acid;
- 6-chloro-8-(2-thienyl)-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;
- 6-chloro-8-(phenylethynyl)-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;
- 10 6-chloro-8-[(4-chlorophenyl)ethynyl]-2-
(trifluoromethyl)-2H-1-benzopyran-3-carboxylic
acid;
- 6-chloro-8-[(4-methoxyphenyl)ethynyl]-2-
(trifluoromethyl)-2H-1-benzopyran-3-carboxylic
15 acid;
- (S)-6-chloro-8-[(4-methoxyphenyl)ethynyl]-2-
(trifluoromethyl)-2H-1-benzopyran-3-carboxylic
acid;
- 6-(phenylethynyl)-2-(trifluoromethyl)-2H-1-
20 benzopyran-3-carboxylic acid;
- 6-chloro-8-(4-chlorophenyl)-2-(trifluoromethyl)-2H-
1-benzopyran-3-carboxylic acid;
- 6-chloro-8-phenyl-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;
- 25 6-(4-bromophenyl)-2-(trifluoromethyl)-2H-1-
benzopyran-3-carboxylic acid;
- 6-chloro-8-(4-methoxyphenyl)-2-trifluoromethyl-2H-1-
benzopyran-3-carboxylic acid; and
- 6-(2,2,2-trifluoro-1-hydroxyethyl)-2-
30 (trifluoromethyl)-2H-1-benzopyran-3-carboxylic
acid.

11. A compound of Claim 2 wherein X is S;
wherein R is carboxyl; wherein R¹ is selected from
35 C₁-C₆-perfluoroalkyl; wherein R² is one or more
radicals independently selected from hydrido, halo,
C₁-C₆-alkyl, phenyl-C₁-C₆-alkyl, phenyl-C₂-C₆-alkynyl,
phenyl-C₂-C₆-alkenyl, C₁-C₆-alkoxy, phenoxy, 5- or

- 6-membered heteroaryloxy, phenyl-C₁-C₆-alkyloxy, 5- or 6-membered heteroaryl-C₁-C₆-alkyloxy, C₁-C₆-haloalkyl, C₁-C₆-haloalkoxy, C₁-C₆-alkylamino, N-phenylamino, N-(phenyl-C₁-C₆-alkyl)amino, N-
- 5 heteroarylamino, N-(heteroaryl-C₁-C₆-alkylamino, nitro, amino, aminosulfonyl, N-alkylaminosulfonyl, N-arylamino, N-heteroarylaminosulfonyl, N-(phenyl-C₁-C₆-alkyl)aminosulfonyl, N-(heteroaryl-C₁-C₆-alkyl)aminosulfonyl, 5- to 8-membered
- 10 heterocyclsulfonyl, C₁-C₆-alkylsulfonyl, optionally substituted phenyl, optionally substituted 5- or 6-membered heteroaryl, phenyl-C₁-C₆-alkylcarbonyl, heteroarylcarbonyl, phenylcarbonyl, aminocarbonyl, and C₁-C₆-alkylcarbonyl; wherein the A ring atoms A¹, A², A³ and A⁴ are independently selected from carbon and nitrogen with the proviso that at least three of A¹, A², A³ and A⁴ are carbon; or an isomer or pharmaceutically acceptable salt thereof.
- 15
- 20 12. A compound of Claim 11 wherein X is S; wherein R is carboxyl; wherein R" is selected from hydrido and ethenyl; wherein R¹ is selected from trifluoromethyl and pentafluoroethyl; wherein R² is one or more radicals independently selected from
- 25 hydrido, chloro, bromo, fluoro, iodo, methyl, tert-butyl, ethenyl, ethynyl, 5-chloro-1-pentynyl, 1-pentynyl, 3,3-dimethyl-1-butynyl, benzyl, phenylethyl, phenyl-ethynyl, 4-chlorophenyl-ethynyl, 4-methoxyphenyl-ethynyl, phenylethenyl, methoxy,
- 30 methylthio, methylsulfinyl, phenyloxy, phenylthio, phenylsulfinyl, pyridyloxy, thienyloxy, furyloxy, phenylmethoxy, methylenedioxy, benzyloxymethyl, trifluoromethyl, difluoromethyl, pentafluoroethyl, trifluoromethoxy, trifluoromethylthio,
- 35 hydroxymethyl, hydroxy-trifluoroethyl, methoxymethyl, hydroxyiminomethyl, N-methylamino, N-phenylamino, N-(benzyl)amino, nitro, cyano, amino, aminosulfonyl, N-methylaminosulfonyl, N-

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- phenylaminosulfonyl, N-furylaminosulfonyl, N-(benzyl)aminosulfonyl, N-(furylmethyl)aminosulfonyl, benzylsulfonyl, phenylethylaminosulfonyl, furylsulfonyl, methylsulfonyl, phenyl, phenyl
5 substituted with one or more radicals selected from chloro, fluoro, bromo, methoxy, methylthio and methylsulfonyl, benzimidazolyl, thienyl, thienyl substituted with chloro, furyl, furyl substituted with chloro, benzylcarbonyl, furylcarbonyl,
10 phenylcarbonyl, aminocarbonyl, formyl, and methylcarbonyl; wherein the A ring atoms A¹, A², A³ and A⁴ are carbon; or an isomer or pharmaceutically acceptable salt thereof.
- 15 13. A compound of Claim/12 selected from compounds, and their isomers and pharmaceutically-acceptable salts, of the group consisting of
6-chloro-2-trifluoromethyl-2H-1-benzothiopyran-3-carboxylic acid;
20 6-methyl-2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;
6,8-dimethyl-2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;
25 6-(1,1-dimethylethyl)-2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;
7-methyl-2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;
6,7-dimethyl-2-(trifluoromethyl)-2H-1-benzothiopyran-30 3-carboxylic acid;
8-methyl-2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;
2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;
35 6-chloro-7-methyl-2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;
7-chloro-2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;

- 6,7-dichloro-2-(trifluoromethyl)-2H-1-benzothiopyran-3-carboxylic acid;
 2-(trifluoromethyl)-6-[(trifluoromethyl)thio]-2H-1-benzopyran-3-carboxylic acid; and
 5 6,8-dichloro-2-trifluoromethyl-2H-1-benzothiopyran-3-carboxylic acid.

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14. A compound of Claim 2 wherein X is NR^a;
 wherein R^a is selected from hydrido, C₁-C₃-alkyl,
 10 phenyl-C₁-C₃-alkyl, acyl and carboxy-C₁-C₃-alkyl;
 wherein R is carboxyl; wherein R¹ is selected from C₁-C₃-perfluoroalkyl; wherein R² is one or more radicals independently selected from hydrido, halo, C₁-C₆-alkyl, phenyl-C₁-C₆-alkyl, phenyl-C₂-C₆-alkynyl,
 15 phenyl-C₂-C₆-alkenyl, C₁-C₆-alkoxy, phenoxy, 5- or 6-membered heteroaryloxy, phenyl-C₁-C₆-alkyloxy, 5- or 6-membered heteroaryl-C₁-C₆-alkyloxy, C₁-C₆-haloalkyl, C₁-C₆-haloalkoxy, C₁-C₆-alkylamino, N-phenylamino, N-(phenyl-C₁-C₆-alkyl)amino, N-heteroarylamino, N-
 20 (heteroaryl-C₁-C₆-alkylamino, nitro, amino, aminosulfonyl, N-alkylaminosulfonyl, N-arylaminosulfonyl, N-heteroarylaminosulfonyl, N-(phenyl-C₁-C₆-alkyl)aminosulfonyl, N-(heteroaryl-C₁-C₆-alkyl)aminosulfonyl, 5- to 8-membered
 25 heterocyclsulfonyl, C₁-C₆-alkylsulfonyl, optionally substituted phenyl, optionally substituted 5- or 6-membered heteroaryl, phenyl-C₁-C₆-alkylcarbonyl, heteroarylcarbonyl, phenylcarbonyl, aminocarbonyl, and C₁-C₆-alkylcarbonyl; wherein the A ring atoms A¹,
 30 A², A³ and A⁴ are independently selected from carbon and nitrogen with the proviso that at least three of A¹, A², A³ and A⁴ are carbon; or an isomer or pharmaceutically acceptable salt thereof.
- 35 15. A compound of Claim 14 wherein X is NR^a;
 wherein R^a is selected from hydrido, methyl, ethyl, (4-trifluoromethyl)benzyl, (4-chloromethyl)benzyl, (4-methoxy)benzyl, (4-cyano)benzyl, and (4-

nitro)benzyl; wherein R is carboxyl; wherein R" is selected from hydrido and ethenyl; wherein R' is selected from trifluoromethyl and pentafluoroethyl; wherein R² is one or more radicals independently
5 selected from hydrido, chloro, bromo, fluoro, iodo, methyl, tert-butyl, ethenyl, ethynyl, 5-chloro-1-pentynyl, 1-pentynyl, 3,3-dimethyl-1-butynyl, benzyl, phenylethyl, phenyl-ethynyl, 4-chlorophenyl-ethynyl, 4-methoxyphenyl-ethynyl, phenylethenyl, methoxy,
10 methylthio, methylsulfinyl, phenoxy, phenylthio, phenylsulfinyl, pyridyloxy, thienyloxy, furyloxy, phenylmethoxy, methylenedioxy, benzyloxymethyl, trifluoromethyl, difluoromethyl, pentafluoroethyl, trifluoromethoxy, trifluoromethylthio, hydroxymethyl,
15 hydroxy-trifluoroethyl, methoxymethyl, hydroxyiminomethyl, N-methylamino, N-phenylamino, N-(benzyl)amino, nitro, cyano, amino, aminosulfonyl, N-methylaminosulfonyl, N-phenylaminosulfonyl, N-furylaminosulfonyl, N-(benzyl)aminosulfonyl, N-
20 (furylmethyl)aminosulfonyl, benzylsulfonyl, phenylethylaminosulfonyl, furylsulfonyl, methylsulfonyl, phenyl, phenyl substituted with one or more radicals selected from chloro, fluoro, bromo, methoxy, methylthio and methylsulfonyl,
25 benzimidazolyl, thienyl, thienyl substituted with chloro, furyl, furyl substituted with chloro, benzylcarbonyl, furylcarbonyl, phenylcarbonyl, aminocarbonyl, formyl, and methylcarbonyl; wherein the A ring atoms A¹, A², A³ and A⁴ are carbon; or an
30 isomer or pharmaceutically acceptable salt thereof.

16. A compound of Claim 15 selected from compounds, and their isomers and pharmaceutically-acceptable salts, of the group consisting of

35 6-chloro-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid;

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17. A compound of Claim 2 wherein X is selected
 from O, S and NR^a; wherein R^a is selected from
 hydrido, C₁-C₆-alkyl, phenyl-C₁-C₆-alkyl, acyl and
 carboxy-C₁-C₆-alkyl; wherein R is selected from
 carboxyl; wherein R¹ is selected from C₁-C₆-alkyl, phenyl-C₁-C₆-alkyl, acyl and carboxy-C₁-C₆-alkyl; and
 (S)-6-chloro-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid.
- 6,8-dichloro-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
- 6,7-difluoro-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
- 5 6-iodo-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
- 6-bromo-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
- 10 1,2-dihydro-6-(trifluoromethoxy)-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
- 6-(trifluoromethyl)-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
- 6-cyano-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
- 15 6-chloro-1,2-dihydro-1-methyl-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
- 6-chloro-1,2-dihydro-2-(trifluoromethyl)-1-[[4-(trifluoromethyl)phenyl]methyl]-3-quinolinecarboxylic acid;
- 20 6-chloro-1-[(4-chlorophenyl)methyl]-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
- 6-chloro-1,2-dihydro-2-(trifluoromethyl)-1-[[4-(methoxy)phenyl]methyl]-3-quinolinecarboxylic acid;
- 25 6-chloro-1-[(4-cyanophenyl)methyl]-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
- 6-chloro-1,2-dihydro-1-[(4-nitrophenyl)methyl]-2-(trifluoromethyl)-3-quinolinecarboxylic acid;
- 6-chloro-1,2-dihydro-1-ethyl-2-(trifluoromethyl)-3-
- 30 quinolinecarboxylic acid; and
 (S)-6-chloro-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid.

17. A compound of Claim 2 wherein X is selected
 from O, S and NR^a; wherein R^a is selected from
 hydrido, C₁-C₆-alkyl, phenyl-C₁-C₆-alkyl, acyl and
 carboxy-C₁-C₆-alkyl; wherein R is selected from
 carboxyl; wherein R¹ is selected from C₁-C₆-alkyl, phenyl-C₁-C₆-alkyl, acyl and carboxy-C₁-C₆-alkyl; and
 (S)-6-chloro-1,2-dihydro-2-(trifluoromethyl)-3-quinolinecarboxylic acid.

perfluoroalkyl; wherein the A ring atoms A¹, A², A³ and A⁴ are independently selected from carbon and nitrogen with the proviso that at least three of A¹, A², A³ and A⁴ are carbon; and wherein R² together with ring A

5 forms a naphthyl or quinolyl radical; or an isomer or pharmaceutically acceptable salt thereof.

18. A compound of Claim 17 wherein X is selected from O, S and NR^a; wherein R^a is selected from

10 hydrido, methyl, ethyl, (4-trifluoromethyl)benzyl, (4-chloromethyl)benzyl, (4-methoxy)benzyl, and (4-cyano)benzyl, (4-nitro)benzyl; wherein R is carboxyl; wherein R^b is selected from hydrido and ethenyl; wherein R^c is selected from trifluoromethyl and

15 pentafluoroethyl; wherein the A ring atoms A¹, A², A³ and A⁴ are independently selected from carbon and nitrogen with the proviso that at least three of A¹, A², A³ and A⁴ are carbon; or wherein R² together with ring A forms a naphthyl, or quinolyl radical; or an

20 isomer or pharmaceutically acceptable salt thereof.

19. A compound of Claim 18 selected from compounds, and their isomers and pharmaceutically-acceptable salts, of the group consisting of

25 2-trifluoromethyl-2H-naphtho[1,2-b]pyran-3-carboxylic acid;

2-trifluoromethyl-3H-naphtho[2,1-b]pyran-3-carboxylic acid;

30 2-trifluoromethyl-2H-naphtho[2,3-b]pyran-3-carboxylic acid;

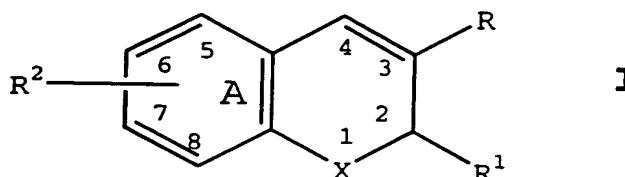
5-(hydroxymethyl)-8-methyl-2-(trifluoromethyl)-2H-pyrano[2,3-c]pyridine-3-carboxylic acid;

6-(trifluoromethyl)-6h-1,3-dioxolo[4,5-

35 g][1]benzopyran-7-carboxylic acid; and

3-(trifluoromethyl)-3H-benzofuro[3,2-f][1]benzopyran-2-carboxylic acid.

20. A compound of Formula I



- 5 wherein X is selected from O or S or NR^a;
- wherein R^a is alkyl;
- wherein R is selected from carboxyl,
aminocarbonyl, alkylsulfonylaminocarbonyl and
10 alkoxycarbonyl;
- wherein R¹ is selected from haloalkyl, alkyl,
aralkyl, cycloalkyl and aryl optionally substituted
with one or more radicals selected from alkylthio,
nitro and alkylsulfonyl; and
- 15 wherein R² is one or more radicals selected from
hydrido, halo, alkyl, aralkyl, alkoxy, aryloxy,
heteroaryloxy, aralkyloxy, heteroaralkyloxy,
haloalkyl, haloalkoxy, alkylamino, arylamino,
aralkylamino, heteroaryl amino, heteroarylalkylamino,
20 nitro, amino, aminosulfonyl, alkylaminosulfonyl,
arylaminosulfonyl, heteroarylaminosulfonyl,
aralkylaminosulfonyl, heteroaralkylaminosulfonyl,
heterocyclosulfonyl, alkylsulfonyl, optionally
substituted aryl, optionally substituted heteroaryl,
25 aralkylcarbonyl, heteroarylcarbonyl, arylcarbonyl,
aminocarbonyl, and alkylcarbonyl;
- or wherein R² together with ring A forms a
naphthyl radical;
- or an isomer or pharmaceutically acceptable salt
30 thereof.

21. Compound of Claim 20 wherein X is oxygen or
sulfur; wherein R is selected from carboxyl, lower
alkyl, lower aralkyl and lower alkoxycarbonyl;

wherein R¹ is selected from lower haloalkyl, lower cycloalkyl and phenyl; and wherein R² is one or more radicals selected from hydrido, halo, lower alkyl, lower alkoxy, lower haloalkyl, lower haloalkoxy,

- 5 lower alkylamino, nitro, amino, aminosulfonyl, lower alkylaminosulfonyl, 5- or 6- membered heteroarylalkylaminosulfonyl, lower aralkylaminosulfonyl, 5- or 6- membered nitrogen containing heterocyclosulfonyl, lower alkylsulfonyl,
- 10 optionally substituted phenyl, lower aralkylcarbonyl, and lower alkylcarbonyl; or wherein R² together with ring A forms a naphthyl radical; or an isomer or pharmaceutically acceptable salt thereof.

- 15 22. Compound of Claim 21 wherein X is oxygen or sulfur; wherein R is carboxyl; wherein R¹ is lower haloalkyl; and wherein R² is one or more radicals selected from hydrido, halo, lower alkyl, lower haloalkyl, lower haloalkoxy, lower alkylamino, amino, aminosulfonyl, lower alkylaminosulfonyl, 5- or 6- membered heteroarylalkylaminosulfonyl, lower aralkylaminosulfonyl, lower alkylsulfonyl, 6- membered nitrogen containing heterocyclosulfonyl, optionally substituted phenyl, lower aralkylcarbonyl, and lower alkylcarbonyl; or wherein R² together with ring A forms a naphthyl radical; or an isomer or pharmaceutically acceptable salt thereof.

23. Compound of Claim 22 wherein R is carboxyl, wherein R¹ is selected from fluoromethyl, chloromethyl, dichloromethyl, trichloromethyl, pentafluoroethyl, heptafluoropropyl, difluoroethyl, difluoropropyl, dichloroethyl, dichloropropyl, difluoromethyl, and trifluoromethyl; and wherein R² is one or more radicals selected from hydrido, chloro, fluoro, bromo, iodo, methyl, ethyl, isopropyl, tert-butyl, butyl, isobutyl, pentyl, hexyl, methoxy, ethoxy, isopropyloxy, tertbutyloxy, trifluoromethyl,

difluoromethyl, trifluoromethoxy, amino, N,N-
dimethylamino, N,N-diethylamino, N-
phenylmethylaminosulfonyl, N-
phenylethylaminosulfonyl, N-(2-
5 furylmethyl)aminosulfonyl, nitro, N,N-
dimethylaminosulfonyl, aminosulfonyl, N-
methylaminosulfonyl, N-ethylsulfonyl, 2,2-
dimethylethylaminosulfonyl, N,N-
dimethylaminosulfonyl, N-(2-
10 methylpropyl)aminosulfonyl, N-morpholinosulfonyl,
methylsulfonyl, benzylcarbonyl, 2,2-
dimethylpropylcarbonyl, phenylacetyl and phenyl; or
wherein R' together with ring A forms a naphthyl
radical; or an isomer or pharmaceutically acceptable
15 salt thereof.

24. Compound of Claim 23 wherein R is carboxyl; wherein R¹ is trifluoromethyl or pentafluorethyl; and wherein R² is one or more radicals selected from
20 hydrido, chloro, fluoro, bromo, iodo, methyl, ethyl, isopropyl, tert-butyl, methoxy, trifluoromethyl, trifluoromethoxy, N-phenylmethylaminosulfonyl, N-phenylethylaminosulfonyl, N-(2-furymethyl)aminosulfonyl, N,N-dimethylaminosulfonyl,
25 N-methylaminosulfonyl, N-(2,2-dimethylethyl)aminosulfonyl, dimethylaminosulfonyl, 2-methylpropylaminosulfonyl, N-morpholinosulfonyl, methylsulfonyl, benzylcarbonyl, and phenyl; or wherein R² together with ring A forms a naphthyl
30 radical; or an isomer or pharmaceutically acceptable salt thereof.

35 25. A compound of Claim 24 selected from
compounds, and their isomers and pharmaceutically-
acceptable salts, of the group consisting of
6-chloro-2-trifluoromethyl-2H-1-benzopyran-3-
carboxylic acid;

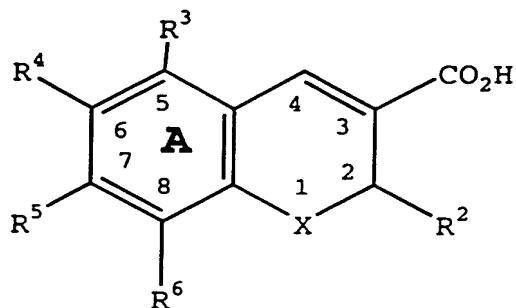
- 6-chloro-7-methyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 8-(1-methylethyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 5 6-chloro-7-(1,1-dimethylethyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-chloro-8-(1-methylethyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 2-trifluoromethyl-3H-naphthopyran-3-carboxylic acid ;
- 10 7-(1,1-dimethylethyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-bromo-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 8-chloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 15 6-trifluoromethoxy-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 5,7-dichloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 20 8-phenyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 7,8-dimethyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6,8-bis(dimethylethyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 25 7-(1-methylethyl)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 7-phenyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 30 6-chloro-7-ethyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;

- 6-chloro-8-ethyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-chloro-7-phenyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 5 6,7-dichloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6,8-dichloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 2-trifluoromethyl-3H-naptho[2,1-b]pyran-3-carboxylic acid;
- 10 6-chloro-8-methyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 8-chloro-6-methyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 15 8-chloro-6-methoxy-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-bromo-8-chloro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 8-bromo-6-fluoro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 20 8-bromo-6-methyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 8-bromo-5-fluoro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 25 6-chloro-8-fluoro-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-bromo-8-methoxy-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-[(phenylmethyl)amino]sulfonyl]-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 30 6-[(dimethylamino)sulfonyl]-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;

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- 6-[(methylamino)sulfonyl]-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-[(4-morpholino)sulfonyl]-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 5 6-[(1,1-dimethylethyl)aminosulfonyl]-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-[(2-methylpropyl)aminosulfonyl]-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-methylsulfonyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 10 8-chloro-6-[(phenylmethyl)amino]sulfonyl]-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-phenylacetyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 15 6,8-dibromo-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 8-chloro-5,6-dimethyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6,8-dichloro-(S)-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 20 6-benzylsulfonyl-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-[[N-(2-furylmethyl)amino]sulfonyl]-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 25 6-[[N-(2-phenylethyl)amino]sulfonyl]-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 6-iodo-2-trifluoromethyl-2H-1-benzopyran-3-carboxylic acid;
- 7-(1,1-dimethylethyl)-2-pentafluoroethyl-2H-1-benzopyran-3-carboxylic acid; and
- 30 6-chloro-2-trifluoromethyl-2H-1-benzothiopyran-3-carboxylic acid.

26. A compound of Formula II



wherein X is O or S;

wherein R² is lower haloalkyl;

5 wherein R³ is selected from hydrido, and halo;

wherein R⁴ is selected from hydrido, halo, lower alkyl, lower haloalkoxy, lower alkoxy, lower

aralkylcarbonyl, lower dialkylaminosulfonyl, lower alkylaminosulfonyl, lower aralkylaminosulfonyl, lower

10 heteroaralkylaminosulfonyl, and 5- or 6- membered nitrogen-containing heterocyclosulfonyl;

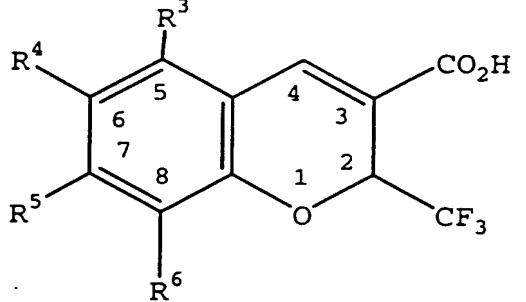
wherein R⁵ is selected from hydrido, lower alkyl, halo, lower alkoxy, and aryl; and

15 wherein R⁶ is selected from hydrido, halo, lower alkyl, lower alkoxy, and aryl;

or an isomer or pharmaceutically acceptable salt thereof.

27. A compound of Formula IIa:

20



wherein R³ is selected from hydrido, lower alkyl, lower hydroxyalkyl, lower alkoxy and halo;

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wherein R⁴ is selected from hydrido, halo, lower alkyl, lower alkylthio, lower haloalkyl, amino, aminosulfonyl, lower alkylsulfonyl, lower alkylsulfinyl, lower alkoxyalkyl, lower

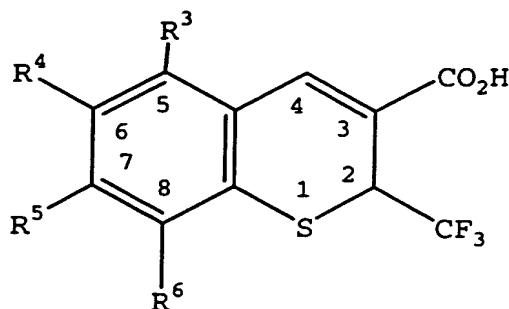
- 5 alkylcarbonyl, formyl, cyano, lower haloalkylthio, substituted or unsubstituted phenylcarbonyl, lower haloalkoxy, lower alkoxy, lower aralkylcarbonyl, lower dialkylaminosulfonyl, lower alkylaminosulfonyl, lower aralkylaminosulfonyl, lower
10 heteroaralkylaminosulfonyl, 5- or 6- membered heteroaryl, lower hydroxyalkyl, optionally substituted phenyl and 5- or 6- membered nitrogen containing heterocyclosulfonyl; wherein R⁵ is selected from hydrido, lower alkyl, halo, lower haloalkyl, lower alkoxy, and phenyl; and wherein R⁶ is selected from hydrido, halo, cyano, hydroxyiminomethyl, lower hydroxyalkyl, lower alkynyl, phenylalkynyl, lower alkyl, lower alkoxy, formyl and phenyl; or an isomer or pharmaceutically acceptable salt
15 thereof.
- 20

28. Compound of Claim 27 wherein R³ is selected from hydrido, and chloro; wherein R⁴ is selected from chloro, methyl, tert-butyl, methylthio,

- 25 trifluoromethyl, difluoromethyl, pentafluoromethyl, trifluoromethylsulfide, trifluoromethoxy, cyano, substituted or unsubstituted phenylcarbonyl, and substituted or unsubstituted phenyl; wherein R⁵ is selected from hydrido, methyl, tert-butyl, chloro, and wherein R⁶ is selected from hydrido, chloro, thienyl, hydroxyiminomethyl, substituted or unsubstituted phenylethynyl, and substituted or unsubstituted phenyl; or an isomer or pharmaceutically acceptable salt thereof.
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29. A compound of Formula IIb:

**IIb**

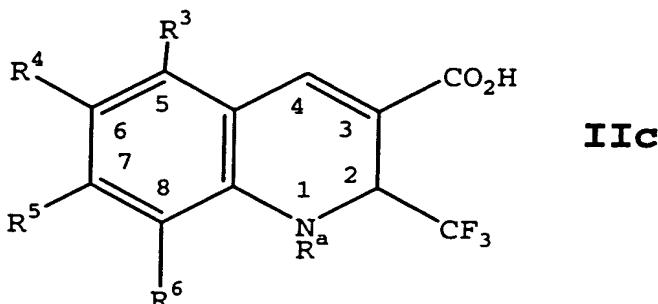
- wherein R³ is selected from hydrido, lower alkyl,
lower hydroxyalkyl, lower alkoxy and halo; wherein R⁴
 5 is selected from hydrido, halo, lower alkyl, lower
alkylthio, lower haloalkyl, amino, aminosulfonyl,
lower alkylsulfonyl, lower alkylsulfinyl, lower
alkoxyalkyl, lower alkylcarbonyl, formyl, cyano,
lower haloalkylthio, substituted or unsubstituted
 10 phenylcarbonyl, lower haloalkoxy, lower alkoxy, lower
aralkylcarbonyl, lower dialkylaminosulfonyl, lower
alkylaminosulfonyl, lower aralkylaminosulfonyl, lower
heteroaralkylaminosulfonyl, 5- or 6- membered
heteroaryl, lower hydroxyalkyl, optionally
 15 substituted phenyl and 5- or 6- membered nitrogen
containing heterocyclosulfonyl; wherein R⁵ is selected
from hydrido, lower alkyl, halo, lower haloalkyl,
lower alkoxy, and phenyl; and wherein R⁶ is selected
 20 from hydrido, halo, cyano, hydroxyiminomethyl, lower
hydroxyalkyl, lower alkynyl, phenylalkynyl, lower
alkyl, lower alkoxy, formyl and phenyl; or an isomer
or pharmaceutically acceptable salt thereof.

30. Compound of Claim 29 wherein R³ is selected
 25 from hydrido, and chloro; wherein R⁴ is selected from
chloro, methyl, tert-butyl, methylthio,
trifluoromethyl, difluoromethyl, pentafluoromethyl,
trifluoromethylsulfide, trifluoromethoxy, cyano,
substituted or unsubstituted phenylcarbonyl, and
 30 substituted or unsubstituted phenyl; wherein R⁵ is
selected from hydrido, methyl, tert-butyl, chloro;

and wherein R⁶ is selected from hydrido, chloro, thienyl, hydroxyiminomethyl, substituted or unsubstituted phenylethynyl, and substituted or unsubstituted phenyl; or an isomer or

- 5 pharmaceutically acceptable salt thereof.

31. A compound of Formula IIc:



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wherein R^a is selected from hydrido and lower aralkyl; wherein R³ is selected from hydrido, lower alkyl, lower hydroxyalkyl, lower alkoxy and halo; wherein R⁴ is selected from hydrido, halo, lower alkyl, lower

- 15 alkylthio, lower haloalkyl, amino, aminosulfonyl, lower alkylsulfonyl, lower alkylsulfinyl, lower alkoxyalkyl, lower alkylcarbonyl, formyl, cyano, lower haloalkylthio, substituted or unsubstituted phenylcarbonyl, lower haloalkoxy, lower alkoxy, lower
20 aralkylcarbonyl, lower dialkylaminosulfonyl, lower alkylaminosulfonyl, lower aralkylaminosulfonyl, lower heteroaralkylaminosulfonyl, 5- or 6- membered heteroaryl, lower hydroxyalkyl, optionally substituted phenyl and 5- or 6- membered nitrogen
25 containing heterocyclosulfonyl; wherein R⁵ is selected from hydrido, lower alkyl, halo, lower haloalkyl, lower alkoxy, and phenyl; and wherein R⁶ is selected from hydrido, halo, cyano, hydroxyiminomethyl, lower hydroxyalkyl, lower alkynyl, phenylalkynyl, lower
30 alkyl, lower alkoxy, formyl and phenyl;

or an isomer or pharmaceutically acceptable salt thereof.

32. Compound of Claim 31 wherein R^a is hydrido;
- 5 wherein R^b is selected from hydrido, and chloro; wherein R^c is selected from chloro, methyl, tert-butyl, methylthio, trifluoromethyl, difluoromethyl, pentafluoromethyl, trifluoromethylsulfide, trifluoromethoxy, cyano, substituted or unsubstituted phenylcarbonyl, and substituted or unsubstituted phenyl; wherein R^d is selected from hydrido, methyl, tert-butyl, chloro; and wherein R^e is selected from hydrido, chloro, thienyl, hydroxyiminomethyl, substituted or unsubstituted phenylethynyl, and substituted or unsubstituted phenyl; or an isomer or pharmaceutically acceptable salt thereof.

33. A method of treating a cyclooxygenase-2 mediated disorder in a subject, said method comprising treating the subject having or susceptible to said disorder with a therapeutically-effective amount of a compound of Claims 1-31; or a pharmaceutically-acceptable salt thereof.

- 25
34. The method of Claim 33 wherein the cyclooxygenase-2 mediated disorder is inflammation.

35. The method of Claim 33 wherein the cyclooxygenase-2 mediated disorder is arthritis.

36. The method of Claim 33 wherein the cyclooxygenase-2 mediated disorder is pain.

- 35
37. The method of Claim 33 wherein the cyclooxygenase-2 mediated disorder is fever.

38. A pharmaceutical composition comprising a therapeutically-effective amount of a compound, said compound selected from a family of compounds of Claims 1-~~31~~; or a pharmaceutically-acceptable salt
5 thereof.

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